L	Hits	Search Text	DB	Time stamp
Number				
4	2303	lipoic adj acid	USPAT;	2004/09/16
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			EPO; JPO	
5	324	(lipoic adj acid ) and platinum	USPAT;	2004/09/16
		(	US-PGPUB;	18:48
			EPO; JPO	
6	1152	"I6" and cancer	USPAT;	2004/09/16
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ľ			EPO; JPO	
7	0	((lipoic adj acid ) and platinum) and cnacer	USPAT:	2004/09/16
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			EPO; JPO	
8	237	((lipoic adj acid ) and platinum) and cancer	USPAT;	2004/09/16
•	231	((iipoic auj acia ) ana piasinam, ana cancer	US-PGPUB;	18:49
			EPO; JPO	
	1	(((lipoic adj acid ) and platinum) and	USPAT:	2004/09/16
9	1	-	US-PGPUB;	18:49
		cancer) and polynuclear	EPO; JPO	

(FILE 'HOME' ENTERED AT 16:04:02 ON 16 SEP 2004)

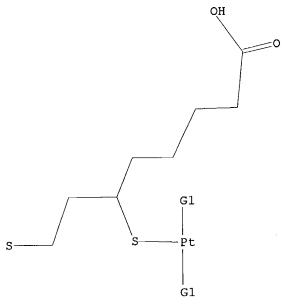
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L1

=> d 11

L1 HAS NO ANSWERS

L1STR



G1 C1,OH,NH3

Structure attributes must be viewed using STN Express query preparation.

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SAMPLE SEARCH INITIATED 16:05:03 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED

0 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

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PROJECTED ITERATIONS:

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FULL SEARCH INITIATED 16:05:09 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 20 TO ITERATE

100.0% PROCESSED

20 ITERATIONS

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SEARCH TIME: 00.00.01

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COST IN U.S. DOLLARS

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FULL ESTIMATED COST ENTRY SESSION 155.84 156.05

FILE 'CAPLUS' ENTERED AT 16:05:22 ON 16 SEP 2004
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FILE COVERS 1907 - 16 Sep 2004 VOL 141 ISS 12 FILE LAST UPDATED: 15 Sep 2004 (20040915/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s lipoic acid and platinum

3162 LIPOIC

3869612 ACID

3114 LIPOIC ACID

(LIPOIC(W)ACID)

182692 PLATINUM

L4 11 LIPOIC ACID AND PLATINUM

=> d 1-11 bib abs

L4 ANSWER 1 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:120684 CAPLUS

DN 140:187383

TI Lipid-drug complexes in reversed liquid and liquid crystalline phases

IN Anderson, David M.

PA Lyotropic Therapeutics, Inc., USA

SO PCT Int. Appl., 51 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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PATENT NO.						D	DATE			APPL:	[CAT]	DATE					
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WO 2004012680 WO 2004012680				A2		20040212			NO 2	003-1		20030806					
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20030806 US 2003-635019 US 2004156816 A1 20040812 PRAI US 2002-401011P 20020806 Р A pharmaceutical is formulated to enable enhanced delivery across membrane barriers, permit solubilization, protect compds. from deactivation by thiol containing compds. in the body, and allow retention of the drug during transport to a desired site of activity. The pharmaceutical includes a complex of two moieties where at least one is pharmaceutically active and is larger than a single atom in size, and the second moiety, when combined with a cationic or anionic counterion forms either a pharmaceutically acceptable anionic or cationic surfactant or a pharmaceutically acceptable salt that has an octanol water partition coefficient of greater than about 100. A composition contained cisplatin in dimethylacetamide and Epikuron 105. ANSWER 2 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN L4

GW, ML, MR, NE, SN, TD, TG

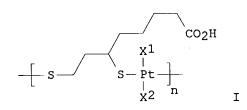
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2004:60263 CAPLUS
ΑN
     140:121563
DN
     Preparation of polynuclear platinum lipoic
ΤI
     acid compounds as anticancer agents
     Lal, Manjari; Palepu, Nagesh
IN
     Sonus Pharmaceuticals, Inc., USA
PA
     PCT Int. Appl., 39 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LА
FAN.CNT 1
                                                APPLICATION NO.
                                                                         DATE
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                                                                         20030716
                                   20040122
                                                WO 2003-US22221
     WO 2004006859
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     WO 2004006859
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              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
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              CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
              NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ,
              GW, ML, MR, NE, SN, TD, TG
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20040513

20020716

A1

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US 2004092585

PRAI US 2002-396299P

GΙ

The preparation is described for cisplatin tocopherol, carboplatin folic acid and polynuclear platinum lipoic acid complexes for use as anticancer agents. Methods are claimed for using the platinum compds., either alone or in combination with at least one addnl. therapeutic agent, in the prophylaxis or treatment of proliferative diseases. Thus, polynuclear platinum lipoic

US 2003-622007

20030716

acid derivative complexes (I; X1 and X2 = chloro, hydroxy or amino, n = 10-20) were prepared and.

= 10-20) were prepared and. ANSWER 3 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN L42003:984085 CAPLUS ΑN 140:349530 DN Functionalized derivatives of  $\beta$ -hydroxydithiocinnamic acids as TIligands. Crystal structure of 4'-hydroxy- $\beta$ -hydroxydithiocinnamic acid Schubert, Karsten; Saumweber, Rupert; Goerls, Helmar; Weigand, Wolfgang ΑU Inst. Anorganische und Anal. Chem., Friedrich-Schiller-Univ. Jena, Jena, CS D-07743, Germany Zeitschrift fuer Anorganische und Allgemeine Chemie (2003), 629(12-13), SO 2091-2096 CODEN: ZAACAB; ISSN: 0044-2313 Wiley-VCH Verlag GmbH & Co. KGaA PBJournal DTGerman LΑ Silyl-protected 4'-hydroxyacetophenone reacted with CS2 and MeI using K AΒ tert-butylate as a base to give silyl-substituted 4'-hydroxy- $\beta$ hydroxydithiocinnamic acid Me ester. Mol. structure of the deprotected ester (I) was determined by x-ray anal. (monoclinic, a 9.3236(2), b 5.4082(1), c 20.4968(5) Å,  $\beta$  100.017(1)°, Z = 4, dc = 1.477, 1869 observed reflections with Fo >  $4\sigma(Fo)$ , R1 = 0.032, wR2 = 0.083). Esterification of I with  $DL-\alpha-$  lipoic acid gave 4'-(1,2-dithiolane-3-pentanoyl)- $\beta$ -hydroxydithiocinnamic acid Me ester (II). Ni(II), Pd(II) and Pt(II) complexes of the ligand II were synthesized. THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 24 ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 4 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN L42002:756358 CAPLUS ΑN DN138:52125 Development of a new assay for the screening of hypochlorous acid TIscavengers based on reversed-phase high-performance liquid chromatography Gatto, Maria Teresa; Firuzi, Omidreza; Agostino, Roberta; Grippa, ΑU Eleonora; Borso, Angela; Spinelli, Francesca; Pavan, Lucia; Petrolati, Marzia; Petrucci, Rita; Marrosu, Giancarlo; Saso, Luciano Dipartimento di Farmacologia delle Sostanze Naturali e Fisiologia Generale CS Universita di Roma "La Sapienza", Rome, 00185, Italy Biomedical Chromatography (2002), 16(6), 404-411 SO CODEN: BICHE2; ISSN: 0269-3879 John Wiley & Sons Ltd. PΒ Journal DTLA English A new assay for the screening of hypochlorite/hypochlorous acid (XOCl) ΑB scavengers, based on the reversed-phase high performance liquid chromatog. anal. of human serum albumin (HSA, 0.2% in 100 mM sodium phosphate, pH 7), before and after oxidation by XOCl (1.6 mM), was developed. XOCl induced a significant decrease of the area under the chromatog. peak of HSA at 280 nm due to the oxidation of the aromatic amino acids tryptophan and tyrosine, as suggested by the literature and by the chromatog. analyses and the

suggested by the literature and by the chromatog. analyses and the electrochem. study performed here. The assay was validated by testing known XOCl scavengers such as ascorbic acid, cysteine, glutathione, S-methylglutathione and  $\alpha$ - lipoic acid and other antioxidants such as carnosine and chlorogenic acid, which inhibited the oxidation of HSA. Quant. activities were calculated using an original formula based on the changes of the area of the albumin peak. Electrochem. data collected here in a homogeneous medium showed that the anodic potentials of the antioxidants tested are less pos. (ascorbic acid, chlorogenic acid and cysteine) or similar ( $\alpha$ - lipoic acid)

compared with those of the aromatic residues (tryptophan and tyrosine) of HSA oxidized by XOCl. However, as expected, carnosine, glutathione and S-methylglutathione were inactive at a glassy-carbon, gold or platinum electrode.

RE.CNT 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L4 ANSWER 5 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2001:866492 CAPLUS
- DN 136:16034
- TI Reactive oxygen species, antioxidants, and the mammalian thioredoxin system
- AU Nordberg, Jonas; Arner, Elias S. J.
- CS Department of Medical Biochemistry and Biophysics, Karolinska Institute, Medical Nobel Institute for Biochemistry, Stockholm, Swed.
- SO Free Radical Biology & Medicine (2001), 31(11), 1287-1312 CODEN: FRBMEH; ISSN: 0891-5849
- PB Elsevier Science Inc.
- DT Journal; General Review
- LA English
- A review. Reactive oxygen species (ROS) are known mediators of AB intracellular signaling cascades. Excessive production of ROS may, however, lead to oxidative stress, loss of cell function, and ultimately apoptosis or necrosis. A balance between oxidant and antioxidant intracellular systems is hence vital for cell function, regulation, and adaptation to diverse growth conditions. Thioredoxin reductase (TrxR) in conjunction with thioredoxin (Trx) is a ubiquitous oxidoreductase system with antioxidant and redox regulatory roles. In mammals, extracellular forms of Trx also have cytokine-like effects. Mammalian TrxR has a highly reactive active site selenocysteine residue resulting in a profound reductive capacity, reducing several substrates in addition to Trx. Due to the reactivity of TrxR, the enzyme is inhibited by many clin. used electrophilic compds. including nitrosoureas, aurothioglucose, platinum compds., and retinoic acid derivs. The properties of TrxR in combination with the functions of Trx position this system at the core of cellular thiol redox control and antioxidant defense. In this review, the authors focus on the reactions of the Trx system with ROS mols. and different cellular antioxidant enzymes. The authors summarize the TrxR-catalyzed regeneration of several antioxidant compds., including ascorbic acid (vitamin C), selenium-containing substances, lipoic acid, and ubiquinone (Q10). The general cellular effects of TrxR inhibition are also discussed. Dinitrohalobenzenes constitute a unique class of immunostimulatory TrxR inhibitors and the authors consider the immunomodulatory effects of dinitrohalobenzene compds. in view of their reactions with the Trx system.
- RE.CNT 299 THERE ARE 299 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L4 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2000:365344 CAPLUS
- DN 133:99527
- TI Dose-dependent protection by **lipoic acid** against cisplatin-induced nephrotoxicity in rats: antioxidant defense system
- AU Somani, Satu M.; Husain, Kazim; Whitworth, Craig; Trammell, Gary L.; Malafa, Mokenge; Rybak, Leonard P.
- CS Department of Pharmacology, Southern Illinois University School of Medicine, Springfield, IL, 62794-9629, USA
- SO Pharmacology & Toxicology (Copenhagen) (2000), 86(5), 234-241 CODEN: PHTOEH; ISSN: 0901-9928
- PB Munksgaard International Publishers Ltd.
- DT Journal
- LA English

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This study was designed to investigate the role of graded doses of
AΒ
     lipoic acid pretreatment against cisplatin-induced
     nephrotoxicity. Male Wistar rats were divided into six groups and treated
     as follows: 1) vehicle (saline) control; 2) cisplatin (16 mg/kg, i.p.); 3)
     lipoic acid (100 mg/kg, i.p.); 4) cisplatin plus
     lipoic acid (25 mg/kg); 5) cisplatin plus lipoic
     acid (50 mg/kg) and 6) cisplatin plus lipoic
     acid (100 mg/kg). Rats were sacrificed three days after
     treatment, and plasma as well as kidneys were isolated and analyzed.
     Plasma creatinine increased (677% of control) following cisplatin
     administration alone which was decreased by lipoic acid
     in a dose-dependent manner. Cisplatin-treated rats showed a depletion of
     renal glutathione (GSH), increased oxidized GSH and decreased GSH/GSH
     oxidized ratio (62%, 166% and 62% of control), resp. which were restored
     with lipoic acid pretreatment. Renal superoxide
     dismutase, catalase, glutathione peroxidase (GSH peroxidase) and
     glutathione reductase activities decreased (62%, 75%, 62% and 80% of
     control), resp., and malondialdehyde content increased (204% of control)
     following cisplatin administration, which were restored with increasing
     doses of lipoic acid. The renal platinum
     concentration increased following cisplatin administration, which was possibly
     decreased by chelation with lipoic acid.
                                              The data
     suggest that the graded doses of lipoic acid
     effectively prevented a decrease in renal antioxidant defense system and
     prevented an increase in lipid peroxidn., platinum content and
     plasma creatinine concns. in a dose-dependent manner.
              THERE ARE 60 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L4
     ANSWER 7 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     2000:291025 CAPLUS
DN
     132:308192
TI
     Preparation of lipoic acid derivatives as antitumor
     agents
IN
     Bingham, Paul M.; Zachar, Zuzana
PΑ
     The Research Foundation of State University of New York, USA
SO
     PCT Int. Appl., 69 pp.
     CODEN: PIXXD2
דת
     Patent
LA
    English
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FAN.	FAN.CNT 1 PATENT NO.						KIND DATE				APPL:			DATE				
ΡI	WO	2000024734				A1 20000504							19991026					
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		6331						2001	1218	1	US 19	999-4		19991026				
		2002				20020	0903		JP 20	000-		19991026						
	US	2002	1072	34		A1		2002	8080	1	US 20	001-9		20010924				
PRAI	US	1998	-105	628P		P	P 19981026											
	US	1999	-427	477		<b>A</b> 3		1999:	1026									

WO 1999-US25140 W 19991026

OS MARPAT 132:308192

This invention relates to the identification of a novel class of therapeutic agents which selectively target and kill tumor cells and certain other types of diseased cells, and to compns. comprising lipoic acid derivs. which poison the pyruvate dehydrogenase complex specifically in such cells. This invention also provides for methods of using therapeutically effective amts. of the lipoic acid derivs. for the treatment of cancer and other diseases. The lipoic acid derivs. described herein have a wide range of preventive and therapeutic applications. In an experiment using mice with melanoma, one group of mice was dosed with 6,8-bisbenzoylmercaptooctanoic acid (at 100 mg/kg) in 10% ethanol; the control group was dosed with 10% ethanol; mice treated with 6,8-bisbenzoylmercaptooctanoic acid had reduced tumor number and mass - less than 30% to 50% as much mass as control.

RE.CNT 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L4 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
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AN 1999:510986 CAPLUS

DN 131:126713

TI Synergistic biocidal activity of ternary complexes of negatively charged biocides, transition metal ions, and neutral chelators

IN Zhu, Benjhan; Schechtman, Svetlana; Chevion, Mordehai

PA Yissum Research Development Company of the Hebrew University of Jerusalem, Israel; Mcinnis, Patricia, G.

SO PCT Int. Appl., 21 pp.

CODEN: PIXXD2

DT Patent

LA English

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rau.	PATENT NO.							DATE			APPL	ICAT	DATE							
PI	WO 9939575						A2 19990			12 WO 1999-US2783							19990209			
			•	BE,		•		DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,		
PRAI		PT, SE 6426093 1998-74039P				B1 P		2002 1998		us 1	999-:	19990209								

AB A biocidal composition composed of a ternary complex of a neg.-charged biocide, a transition metal ion and chelator has synergistic biocidal effects, as compared with a composition of each of the components alone. The neg. charged biocide is chlorophenol, nitrophenol, chlorophenoxyacetic acid, etc. The chelator is preferably a neutral or pos.-charged chelator, such as 1,10-phenanthroline, 2,2'-bipyridyl, 2,2'-biquinoline,etc. The ternary complex may be used for killing or inhibiting the growth of living cells, bacteria, fungi, etc..

- L4 ANSWER 9 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 1992:520336 CAPLUS
- DN 117:120336
- TI Preparation and characterization of modified electrode by self-assembling ferrocene derivative
- AU Tsutsumi, Hiromori; Furumoto, Shozo; Morita, Masayuki; Matsuda, Yoshiharu
- CS Fac. Eng., Yamaguchi Univ., Ube, 755, Japan
- SO Journal of the Electrochemical Society (1992), 139(6), 1522-5 CODEN: JESOAN; ISSN: 0013-4651
- DT Journal
- LA English
- AB Ferrocenylmethyl-1,2-dithiolane-3-pentanoate, which can be used to modify a Au electrode surface, was prepared by a condensation reaction with

hydroxymethylferrocene and 1,2-dithiolane-3-pentanoic acid (D,L- $\alpha$ -lipoic acid). The condensation product has a 1,2-dithiolane ring which adheres to Au surfaces and a ferrocenyl group which is a redox site. The ferrocene rings on the modified electrode were electroactive in both MeCN and aqueous media.

- L4 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 1992:457576 CAPLUS
- DN 117:57576
- TI Preparation and characterization of redox active molecular assemblies on microelectrode arrays
- AU Frisbie, C. D.; Fritsch-Faules, I.; Wollman, E. W.; Wrighton, M. S.
- CS Dep. Chem., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA
- SO Thin Solid Films (1992), 210-211(1-2), 341-7 CODEN: THSFAP; ISSN: 0040-6090
- DT Journal
- LA English
- AΒ Microelectrode arrays, consisting of 6 or 8 individually addressable Au or Pt microelectrodes .apprx.2 µm wide, 50 µm long, and 0.1 µm thick separated by .apprx.2  $\mu m$  on a Si3N4 substrate, can be modified by immersion into a solution containing mols. having thiol, dithiocarbamate, or disulfide functional groups. The functional groups yield selective modification of the Au or Pt, not the Si3N4, with .apprx.1 monolayer of mol. reagents. Electrochem. and Auger electron spectroscopy (AES) data are summarized to illustrate that the dithiocarbamate functional group can be used to link redox active mols. to Au or Pt surfaces. Results are presented to illustrate that secondary ion mass spectrometry (SIMS) can be used to characterize organic monolayers on the microelectrodes. Preliminary findings are presented showing that the esters of lipoic acid, a 5-membered cyclic disulfide, will selectively modify Au surfaces vs. Si3N4, and that the cyclic disulfide will kinetically compete with a linear disulfide for sites on a Au surfaces. In a competition with the linear disulfide, the cyclic disulfide is at least 10-fold more reactive towards Au. Overall, the studies define classes of expts. needed to develop rational approaches to the modification of surfaces using spontaneous self-assembly methods by taking advantage of selective surface coordination chemical of mols. having appropriate functional groups.
- L4 ANSWER 11 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 1991:554162 CAPLUS
- DN 115:154162
- TI Measurement of biological thiols and disulfides by high-performance liquid chromatography and electrochemical detection of silver mercaptide formation
- AU Kuninori, Toyo; Nishiyama, Junko
- CS Osaka Women's Univ., Sakai, 590, Japan
- SO Analytical Biochemistry (1991), 197(1), 19-24 CODEN: ANBCA2; ISSN: 0003-2697
- DT Journal
- LA English
- AB A rapid and sensitive method is described for the measurement of picomole levels of the biol. thiols glutathione, cysteine, penicillamine, cysteamine, and ergothioneine by a combination of HPLC and electrochem. detection (ECD). The compds. were separated isocratically on a reversed-phase C18 column by ion-pair chromatog. with a mobile phase containing 5 mM acetic acid and 2.5 mM sodium 1-octanesulfonate. After chromatog. separation, the eluate was combined with silver nitrate dissolved in ammonium nitrate buffer at pH 10.5. A platinum disc electrode was used at -0.1 V vs. Ag/AgCl to detect the amount of silver ions that had been consumed by the reaction with thiols. For measurement of disulfide, S-sulfonation with sodium sulfite or electroredn. was used to cleave the disulfide, and the thiol anions produced were detected by HPLC-ECD as for the reduced

forms. The method was used to assay thiols and disulfides in biol. materials.